Serial No.: N/A Filed: Herewith
Title: Novel Cark Protein and Nucleic Acid Molecules and Uses

Therefor

Atty/Agent: Jean M. Silveri

Attorney Docket No.: MPI98-105P1RCP2DV1M

1/35

NUCLEIC ACID SEQUENCE

GTCGACCCACGCGTCCGGCCCTGGAGAAAGGAAACTTATAATAAATG GGAAATTATAAATCTAGACCAACCCAAACTTGTACTGATGAATGGAAGAA AAAAGTCAGTGAATCATATGTTATCACAATAGAAGATTAGAAGATGACC TGCAGATCAAGGAAAAAGAACTGACAGAACTAAGGAATATATTTGGCTCT GATGAAGCCTTCAGTAAAGTCAATTTAAATTACCGCACTGAAAATGGGCT GTCTCTACTTCATTTATGTTGCATTTGTGGAGGCAAGAAATCACATATTC GAACTCTTATGTTGAAAGGGCTCCGCCCATCTCGACTGACAAGAAATGGA TTTACAGCCTTGCATTTAGCAGTTTACAAGGATAATGCAGAATTGATCAC TTCTCTGCTTCACAGTGGAGCTGATATACAGCAGGTTGGATACGGTGGCC TCACTGCCCTCCATATTGCTACAATAGCTGGCCACCTAGAGGCTGCTGAT GTGCTGTTGCAACATGGAGCTAATGTCAATATTCAAGATGCAGTTTTTTT CACTCCATTGCATATTGCAGCGTACTATGGACATGAACAGGTAACTCGCC TTCTTTTGAAATTTGGTGCTGATGTAAATGTAAGTGGTGAAGTTGGAGAT AGACCCCTCCACCTAGCATCTGCAAAAGGATTCTTGAATATTGCAAAACT CTTGATGGAAGAAGCAGCAAAGCAGATGTGAATGCTCAAGATAATGAAG ACCATGTCCCACTCCATTTCTGTTCTCGATTTGGACACCATGATATAGTT AAGTATCTGCTGCAAAGTGATTTGGAAGTTCAACCTCATGTTGTTAATAT CTATGGAGATACCCCCTTACACCTGGCATGCTACAATGGCAAATTTGAAG TTGCCAAGGAAATCATCCAAATATCAGGAACAGAAAGTCTGACTAAGGAA **AACATCTTCAGTGAAACAGCTTTTCATAGTGCTTGTACCTATGGCAAGAG** CATTGACCTAGTCAAATTTCTTCTTGATCAGAATGTCATAAACATCAACC CACATTCGCCTGGTTCAGTTCTTACTGGATAATGGAGCTGATATGAATCT AGTGGCTTGTGATCCCAGCAGGTCTAGTGGTGAAAAAGATGAGCAGACAT GTTTGATGTGGGCTTATGAAAAAGGGCATGATGCCATTGTCACACTCCTG **AAGCATTATAAGAGACCACAAGATGAATTGCCCTGTAATGAATATTCTCA** GCCTGGAGGAGATGGCTCCTATGTGTCTGTTCCATCACCCTTGGGGAAGA TTAAAAGCATGACAAAAGAGAAGGCAGATATTCTCCTCCTAAGAGCTGGA **TTGCCTTCACATTTCCATCTTCAGCTCTCAGAAATTGAGTTCCATGAGAT** TATTGGCTCAGGTTCTTTTGGGAAAGTATATAAAGGACGATGCAGAAATA **AAATAGTGGCTATAAAACGTTATCGAGCCAATACCTACTGCTCCAAGTCA** GATGTGGATATGTTTTGCCGAGAGGTGTCCATTCTCTGCCAGCTCAATCA AGATGTTGCCAAAGGCATGGAGTACCTTCACAACCTGACACAGCCAATTA TACATCGTGACTTGAACAGTCACAATATTCTTCTCTATGAGGATGGGCAT GCTGTGGTGGCAGATTTTGGAGAATCAAGATTTCTACAGTCTCTGGATGA AGACAACATGACAAAACAACCTGGGAACCTCCGTTGGATGGCTCCTGAGG TGTTCACGCAGTGCACTCGGTACACCATCAAAGCAGATGTCTTCAGCTAT GCTCTGTGTCTGTGGGAAATTCTCACTGGCGAAATTCCATTCGCTCATCT CAAGCCAGCGGCTGCGGCAGCAGACATGGCTTACCACCACATCAGACCTC CCATTGGCTATTCCATTCCCAAGCCCATATCATCTCTGCTGATACGAGGG TGGAACGCATGTCCTGAAGGAAGACCCGAATTTTCTGAAGTTGTCATGAA **GTTAGAAGAGTGTCTCTGCAACATTGAGCTGATGTCTCCTGCATCAAGTA** ACAGCAGTGGGTCTCTCACCTTCTTCTTCTTCTGATTGCCTGGTGAAC CGGGGAGGACCTGGCCGGAGTCATGTGGCAGCATTAAGAAGTCGTTTCGA ATTGGAATATGCTCTAAATGCAAGGTCCTATGCTGCTTTGTCCCAAAGTG CTGGACAATATTCCTCTCAAGGTCTGTCTTTGGAGGAGATGAAAAGAAGT CTTCAATACACACCCATTGACAAATATGGCTATGTATCCGATCCCATGAG CTCAATGCATTTCATTCTTGCCGAAATAGTAGCAGCTTTGAGGACAGCA GCTGACAGCATTCGGCGTATACCTAAGGAGAGTTTTTTCCCCGAACTGAC AGCAACGATTCCAACCACGGCAAGCTGGCTTCCAACTATAACATTTTACT CTCAAAGGTCTCCTTAAATTGGGCTTGTTTTTACTTGTCCTATTTAATTC

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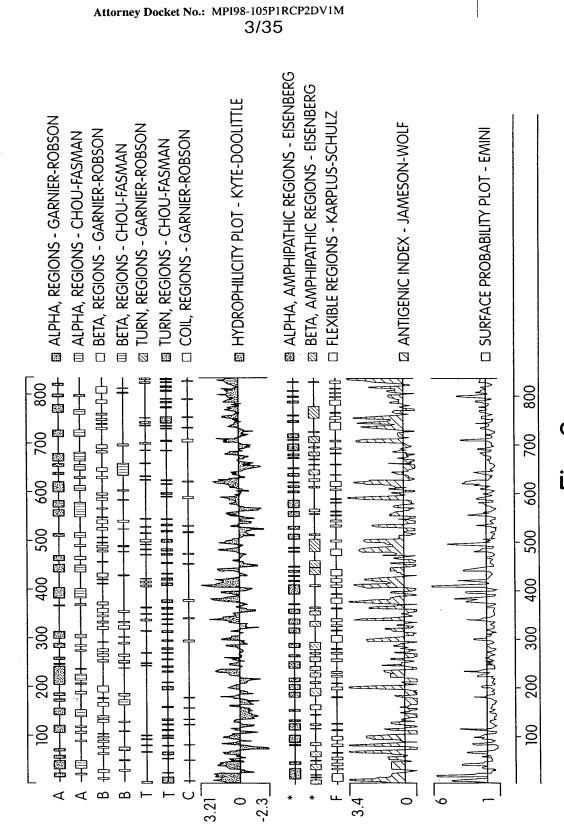
Attorney Docket No.: MPI98-105P1RCP2DV1M

2/35

AMINO ACID SEQUENCE

MGNYKSRPTQTCTDEWKKKVSESYVITIERLEDDLQIKEKELTELRNIFGSDEAFSKVNL
NYRTENGLSLLHLCCICGGKKSHIRTLMLKGLRPSRLTRNGFTALHLAVYKDNAELITSL
LHSGADIQQVGYGGLTALHIATIAGHLEAADVLLQHGANVNIQDAVFFTPLHIAAYYGHE
QVTRLLLKFGADVNVSGEVGDRPLHLASAKGFLNIAKLLMEEGSKADVNAQDVEDHVPLH
FCSRFGHHDIVKYLLQSDLEVQPHVVNIYGDTPLHLACYNGKFEVAKEIIQISGTESLTK
ENIFSETAFHSACTYGKSIDLVKFLLDQNVININHQGRDGHTGLHSACYHGHIRLVQFLL
DNGADMNLVACDPSRSSGEKDEQTCLMWAYEKGHDAIVTLLKHYKRPQDELPCNEYSQPG
GDGSYVSVPSPLGKIKSMTKEKADILLLRAGLPSHFHLQLSEIEFHEIIGSGSFGKVYKG
RCRNKIVAIKRYRANTYCSKSDVDMFCREVSILCQLNHPCVIQFVGACLNDPSQFAIVTQ
YISGGSLFSLLHEQKRILDLQSKLIIAVDVAKGMEYLHNLTQPIIHRDLNSHNILLYEDG
HAVVADFGESRFLQSLDEDNMTKQPGNLRWMAPEVFTQCTRYTIKADVFSYALCLWEILT
GEIPFAHLKPAAAAADMAYHHIRPPIGYSIPKPISSLLIRGWNACPEGRPEFSEVVMKLE
ECLCNIELMSPASSNSSGSLSPSSSSDCLVNRGGPGRSHVAALRSRFELEYALNARSYAA
LSQSAGQYSSQGLSLEEMKRSLQYTPIDKYGYVSDPMSSMHFHSCRNSSSFEDS

Fig. 1B



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Title: Novel Cark Protein and Nucleic Acid Molecules and Uses

Applicants: Jeyaseelan Raju

Atty/Agent: Jean M. Silveri

Serial No.: N/A

Fig. 2

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Therefor

Atty/Agent: Jean M. Silveri

Attorney Docket No.: MPI98-105P1RCP2DV1M

	EEI	Majority
1	GNYKSRPTOTCTDEWKKKVSESYVITTER	CARP2/prot
1	E K K S E E	A.thaliana kinase2
1	S G L C S C S S S G S E G E E G F D A	A.thaliana3
1 1	RTF SDEI.KKKTGFGVGVVPGP	Arabidopsis thaliana c.elegans kinase
ī	RTFSDELKKKISEGYSVVRSR	D.discoideum (A35670)
1	T E T T R M E E D - Q I S C - S	D.Discoideum (U01064)
1	EEEEGAVAKEWGTTPAGPVWTAVFD	H.sapiens (Z48615)
1 1	E E E E G A V A K E W G T T P A G P V W T A V F D E H I Q G A W - K T I S N G F	Homo sapiens (Z48615)
1	EHIQGAW-KTISNGF	Human raf1(W13107) Human Raflkinase(R98215)
ī	GEDGNSWIRRTNFS-HTVCHR	Soybean kinase(M67449)
		-
	L	Majority
	40 50 60 EDDLQIKEKELTELRN	
31	EDDLQIKEKELTELRN	CARP2/prot
8		A.thaliana kinase2
6	N P F R L	A.thaliana3
73 73	Y	Arabidopsis thaliana c.elegans kinase
1		D.discoideum (A35670)
1 6	D V Y A Y A F V L W E	D.Discoideum (U01064)
27	EAAGDEELTLRRGD-RVOVISODC	H.sapiens (Z48615)
27	E A A G D E E L T L R R G D - R V Q V L S Q D C	Homo sapiens (Z48615)
16	GFKDAVFDGS	Human raf1(W13107)
22	DPARLGSI	Human Raflkinase(R98215) Soybean kinase(M67449)
	r with dollar and a second a second and a second a second and a second a second and	Soybean Kinase(Mo/443)
		Majority
	70 80 90	
48	FGSDEAFSKVNLNYRTENGLSLLHLCCIC	CARP2/prot
8		A.thaliana kinase2
12	RW	A.thaliana3
39	AQRKLGW	Arabidopsis thaliana
53 1	MĈPENAELKSTQLLSLFHIICA-	c.elegans kinase
	LTSHLPFRK	D.discoideum (A35670) D.Discoideum (U01064)
	VSGDEGWWTGQL PSGRVGVFPSNYVA	H.sapiens (Z48615)
51	VSGDEGWWTGOL PSGRVGVFPSNYVA	Homo sapiens (Z48615)
26		Human raf1(W13107)
25		Human Raflkinase (R98215)
34	A95AV9442	Soybean kinase(M67449)

Fig. 3A

Filed: Herewith

Title: Novel Cark Protein and Nucleic Acid Molecules and Uses

Therefor

Atty/Agent: Jean M. Silveri

Attorney Docket No.: MPI98-105P1RCP2DV1M

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78 8	G -	G F	K K	S -	H -	I F	T	L -	-	- I	M L	K -	G -	L -	R -	P	<u> </u>	_			- - -	-	s -	R L	- L20 T	CARP2/prot A.thaliana kinase2
47	-			_	_		-	_	_			_	-	_	_							_	_	R -	_	A.thaliana3 Arabidopsis thaliana
76	G	H S	B	S	Q	PE	K	L	Q	F I	LΙ	D	N	L	P	K I	E	S	S I	[]	Ľ	I	S	S Q	S	c.elegans kinase
38	_	- E	N	D	Ī	 s v	- ' A	- . A	- K	v i	A Y	E	- N	L	- R	- : P :	- K :	- · I :	 P 1	 r s	- } -	C	- P	 L -	_	D.discoideum (A35670) D.Discoideum (U01064)
78	P	G A	P	A	A :	P A	G	L	Q		- L	P	Q	E	Ι	P	F	H I	E I	<u>.</u> (L	E	E	ΙI	G	H.sapiens (Z48615)
30	P	G A	. P	Q	A .	P A 	. G	<u>-</u>	Õ	- ·	3 Y	. O	Q R	E	I R	P. A.	ri Si	н. В 1	D (. (3 P	l l	E	E	II PS	G K	Homo sapiens (Z48615) Human raf1(W13107)
30	P	T]	: V	Q	-		-	-	Q	F(3 Y	, O	R	-	R.	A :	SI	D 1	D (3 F	L	T	D	P S	K	Human Raflkinase(R98215)
44																								A -		Soybean kinase(M67449)
	_		-	-	_		_	-	-	- 1	R G	A	R	T	L	V I	K 1	K I	R I	7	A	_	-	D D 1	S	Majority
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47	-		-	-	G (QF	S	T	٧	W I	LΑ	Y	D	Т	L '	r	S :	r	YV	I A	L	K	I	ОК	S	Arabidopsis thaliana
106 1																								V D		c.elegans kinase D.discoideum (A35670)
63																										D.Discoideum (U01064)
106	V	GG	F	G	K '	V Y	R	A	L	WE	R G	E	E	٧.	A '	V I	κ.			- A	A	R	L	DΡ	E	H.sapiens (Z48615)
106	V	GG	F	G	K '	VΥ	R	A	L	W	R G	E	E	V .	A '	V I	K :			- A	A	R	L	D P	E	Homo sapiens (Z48615)
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70	_		_	-	_		=	-	-	FF	₹E	Ä	R	-	Ľ.	E (2 1	KI	RI	S	T	-	-	PN	P	Soybean kinase(M67449)
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57																										A.thaliana kinase2 A.thaliana3
72	Α	QQ	F	Α	Q i	A A	_	-	-			-	_	-	_ ,						_	_	_		_	Arabidopsis thaliana
136 1	D -	Q S	G	R	H]	L L 	P	A	L:	ΗI	À	A	M	I	G]	0 8	S I	E 1	I I	T	'I	L	L	N S	G	c.elegans kinase
78																									_	D.discoideum (A35670) D.Discoideum (U01064)
: 133	K	D P	A	٧	T	ΑE	Q	V	C	Q E	A	R	L	F	G I	A I	<u>د</u> () I	I P	N	I	I	A :	LR	G	H.sapiens (Z48615)
133	K :	D P	A	V	T 1	ΑE	Q	V	C	Q F	A	R	L	F	3 1	A I	, Ç) [IP	N	Į	I	A :	L R	G	Homo sapiens (Z48615)
82 82	_		_	-		- -	_	_	_		. <u>-</u>	_	_	_				- <u>1</u> - I	M د	ı K I K	. А : А	L	K.	v K V R	G G	Human raf1(W13107) Human Raf1kinase(R98215)
86	R :	RE	K	R	I	4 G	K	L	L	N F	D	S	R	E '	r ·					-	_	-	-		-	Soybean kinase(M67449)

Fig. 3B

Applicants: Jeyaseelan Raju Serial No.: N/A Filed: Herewith Title: Novel Cark Protein and Nucleic Acid Molecules and Uses

Therefor

Atty/Agent: Jean M. Silveri

Attorney Docket No.: MPI98-105P1RCP2DV1M

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•			•				1	90)						2	00)							210)
30 69 80 166	A N			- V - P P P	T I P I P I C E C	F	V C C A A	- H - L L V V	F T L - F R	- A L L	- LL LL	H II H II H II H II	E I A M M M M M M	- E T - E E	- F Y - Y Y	- L F - A : A : - :	SG R (R)	- · · · · · · · · · · · · · · · · · · ·	A A A A B B B B B B B B B B B B	DIA	- G V S S A A	D R R R R R	- L T : - V : L : L : L	L I L A L A D W D W	A.thaliana3 Arabidopsis thaliana c.elegans kinase D.discoideum (A35670) D.Discoideum (U01064)
-		-	-	-		-	_	-		_	_			-	_	_				_	L	_	-		Majority
							2	20)						2	30								240	ı
95 188 188 115 115		S R R R R P D	- - A - V V A A	D N - P P A A -	V N	I V I	S G	G G E E -	E V L - L E L	G N V	D D D N N	R I K 7 R I 		H - CH - QQ	L - V L - V V	A : V ! A ! A ! A !	S ; R ; R ; R ; R ; R ; R ; R ; R ; R ;	A H 	G G G G G G G G G G G G G G G G G G G	F F H L - YYF F F F F F F F F F F F F F F F F F	L - FT - L L L L -	N - KS - HIDDI	I I I I I I I I I I I I I I I I I I I	AK SGIK K DA K	CARP2/prot A.thaliana kinase2 A.thaliana3 Arabidopsis thaliana c.elegans kinase D.discoideum (A35670) D.Discoideum (U01064) H.sapiens (Z48615) Homo sapiens (Z48615) Human raf1(W13107) Human Raf1kinase(R98215) Soybean kinase(M67449)
-		-	L	-	_	· -	-	-		_	-	ւ -	-	-	-				K	L	A	-		 270	Majority
218 30 69 112 226	P V P V - V	PPP	L L - L I I L L	L	M	E	G PLLFF	S - A - G K K A A A	K A K A - I S I R K R K	D D - RNNTT	V P - S I I F F	N	A Q A A I I L L L L L L L L L L L L L L L L	D - D - D E E	N R D - Q A A	E Q	D II 	H V E F N Q N H N H H N H	PH GA EINK	LL QL - LL	HS-HH-QAAAA	F (L (Y	C : C : C : C : C : C : C : C : C : C :	S R	CARP2/prot A.thaliana kinase2 A.thaliana3 Arabidopsis thaliana

Applicants: Jeyaseelan Raju Serial No.: N/A Filed: Herewith

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Attorney Docket No.: MPI98-105P1RCP2DV1M

```
300
245 FGHHDIVKYLLQSDLEVQPHVV - - NIYG - - CARP2/prot
A.thaliana kinase2
                                  A.thaliana3
119 ---- MVLEFL---------- Arabidopsis thaliana
251 SGSLVILNMLIKQVRGTNDRICARNLYG-- c.elegans kinase
1 ----- D.discoideum (A35670)
119 ---- D.Discoideum (U01064)
                                  D.Discoideum (U01064)
241 DTVLKITDFGLAREWHKTTKMSAAGTYAWM H.sapiens (Z48615)
241 DTVLKITDFGLAREWHKTTKMSAAGTYAWM Homo sapiens (Z48615)
125 ----- Soybean kinase (M67449)
  -----L---- K--VWS-ILQ-LL---- Majority
           310
                      320
                                330
271 - DTPLHLACYNGKFEVAKEIIQISGTESLT CARP2/prot
125 GDSLLRLIRYNQ-------- Arabidopsis thaliana
279 - DTALHLSCYSGRLDIVKSILDSSPTNIVN c.elegans kinase
    -----
                                  D.discoideum (A35670)
119 ----- KKERFNEITEFLRGKK-- D.Discoideum (U01064)
271 APEVIRLSLFSKSSDVWS--FGVLLWELLT H.sapiens (Z48615)
271 APEVIRLSLFSKSSDVWS - - FGVLLWELLT Homo sapiens (Z48615)
174 EHCSTKVP---TMCVDWSNIRQLLL---- Human raf1(W13107)
174 EHCSTKVP---TMCVDWSNIRQLLL---- Human Raf1kinase(R98215)
125 - - - - - - - - KDSAWTKLLDNGGGKITA Soybean kinase(M67449)
                             ---- Majority
           340
                      350
                                360
300 KENIFSETAFHSACTYGKSIDLVKFLLDQN CARP2/prot
48 PKA----- A.thaliana kinase2
73 KGE----- A.thaliana3
137 - - - YKGLKLN - - - - - KVREICRCIL - - - Arabidopsis thaliana
308 MENVFSETPLHAACTGGKSIELVSFLMKYP c.elegans kinase
  ----- D.discoideum (A35670)
135 - - - - - - - - - -
                                  D.Discoideum (U01064)
299 GEVPYREID - ALAVAYG - - VAMNKLTLPIP H.sapiens (Z48615)
299 GEVPYREID-ALAVAYG--VAMNKLTLPIP Homo sapiens (Z48615)
196 - - - FPNSTI - - - GDSGVPALPSLTMR - - Human raf1(W13107)
196 ----FPNSTI----GDSGVPALPSLTMR-- Human Raf1kinase(R98215)
143 V E T - - - - - - - - - - - - - - - Soybean kinase (M67449)
```

Fig. 3D

Applicants: Jeyaseelan Raju Serial No.: N/A Filed: Herewith Title: Novel Cark 'rotein and Nucleic Acid Molecules and Uses

Therefor

Atty/Agent: Jean M. Silveri

Attorney Docket No.: MPI98-105P1RCP2DV1M

		Wadanita
370	380	390 majority
		
	GHTGLHSACY	
51		A. Challana Kinasez
154	TGL	A. CHAIIAHAS
336 GAD BMAUGUD (GRAFIAGYCA	Arabidopsis thaliana
1	GHTALHSACY	HGHLRI c.elegans kinase
135		D.GIBCOIGEMM (ASSO/O)
	LLEECWDPDP	
326 STCPEPFAR	LIEECWDPDP	HGRPDF Home comices (740C1E)
216		RM Human raf1(W13107)
216		RM Human Raf1kinase(R98215)
146		Soybean kinase (M67449)
<u>-</u>		_ ,
<u>L</u>		E Majority
400	410	420
356 VOFL LDNG	A D M N L V A	CARP2/prot
51		KEE- A.thaliana kinase2
76		REE - A thaliana?
157 - DYLHR		luchidonain bhaliana
364 VQYL LENGI	A D Q S L A S R A F E G - G R P F G G	ALRQQ - c.elegans kinase
1	RPFGG	D.discoideum (A35670)
		D.DISCOIDEUM (001004)
	EQSALFQMPLESFH	Dupicino (210013)
210 BEGAGBMDAG	EQSALFQMPLESFH SQHRYSTPHAFTFN	buplonb (210025)
210 RESVERMEVS	SQHRYSTPHAFTFN	
146		
		Dolinean vinage (mo. 443)
	440	Majority
430	440	450
201	SRSSGEK	
371	S K S S G E K	
70		iii ciida talla hallasca
162 E L G M T H = = = = -	SDLKPENIL	n. charranas
390 A G P (GTNRPSKVASAIMA	
	· ·	C.CICGUID RINGSC
135		D.discoideum (M35670)
	LRTKEKELRSREEE	
381 KLEIQHMFDDI	LRTKEKELRSREEE	L L R A Homo sapiens (748615)
248 E G S I	LSQRQRSTSTPNVH	MVSTTL Human raf1(W13107)
	LSQRQRSTSTPNVH	MVSTTL Human Raf1kinase(R98215)
149		Soybean kinase(M67449)

Fig. 3E

Applicants: Jeyaseelan Raju Serial No.: N/A Filed: Herewith Title: Novel Cark Protein and Nucleic Acid Molecules and Uses

Therefor

Atty/Agent: Jean M. Silveri

Attorney Docket No.: MPI98-105P1RCP2DV1M

		_	_	_	_	_	_		_		_	_	_	_	_	. .				_	W	E	L	D	_	_	-			Majority
					•					46	0								47	0								48	0	Majority
4: 6: 1: 4: 4: 2:	13 35 09 09 71 71	P P P	S - A A V V	S - QQDD	N - EESS	A - Q Q	S - R R R R -	- · · · · · · · · · · · · · · · · · · ·			V - Q Q A A -		L - R R R R -	D R R S S -	D:	E C C C C C C C C C C C C C C C C C C C			V - AAAAA	M E E P P -	WWW-WW-RRSSW	AET - AEEEEAAN	YIA-YTIMMLLV	EDD - EQKDDSSD	K R S - I I S S -	G - V I I I I I I I I I I I I I I I I I I	H - RH - P P	D A		CARP2/prot A.thaliana kinase2 A.thaliana3 Arabidopsis thaliana c.elegans kinase D.discoideum (A35670) D.Discoideum (U01064) H.sapiens (Z48615) Homo sapiens (Z48615) Human raf1(W13107) Human Raf1kinase(R98215) Soybean kinase(M67449)
		L	_	-	_	-	-				_	_	-	_	-	<u>-</u>		-	· -	_	-	_	_	-	_	-	_	 5‡	-	Majority
										49	0								5 Ò	0								51	0	
8: 1: 4: 1: 4: 3:	3 92 43 2 38 38 38 00		TV - HHSS	- PA LLPP	- L L 	L - L L L - M M	K K S - C C	H ! !		K	- - - - - E E - -	R KRR - KK	P P T P	Q - EVP - RR	D	E 1	LH	P C	N - GS - GG TT -	E - A E - N N G G -	Y Y - F W W -	S S KKSS -	Q R R Q Q -	P - S AAPP -	G - T I	— — — — — — — — — — — — — — — — — — —		L R		CARP2/prot A.thaliana kinase2 A.thaliana3 Arabidopsis thaliana c.elegans kinase D.discoideum (A35670) D.Discoideum (U01064) H.sapiens (Z48615) Homo sapiens (Z48615) Human raf1(W13107) Human Raf1kinase(R98215) Soybean kinase(M67449)
		_	_	_	_	_	-		- !	52	 n	_	_	_	-		_	_	- 53	_ n		-	_	_	L		- !	R -	^	Majority
8 2 4 2 1 4 3	8 3 14 66 1 38 68 10	- EG - EE	- E - G G -	- - - - - - - - - - - -	S S S	- Y - H H	V T I I T	S 1 P 1 P 1 S 1 P 1			P PP - GGQQ	L - MP - FFRR	G GP - EEEE	K R Q - H H R R	I i i	X :	S M S I K I V S	T I	K - R - VVTT	E D QQQQ	K K A A E E	A A S S K K	D - K D - P P N N	I KL TTKK	L: 	L I	L I	RA RA RS RK RK RG		CARP2/prot A.thaliana kinase2 A.thaliana3 Arabidopsis thaliana c.elegans kinase D.discoideum (A35670) D.Discoideum (U01064) H.sapiens (Z48615) Homo sapiens (Z48615) Human raf1(W13107) Human Raf1kinase(R98215) Soybean kinase(M67449)

Fig. 3F

Serial No.: N/A Filed: Herewith

Title: Novel Cark Protein and Nucleic Acid Molecules and Uses

Therefor

Atty/Agent: Jean M. Silveri

Attorney Docket No.: MPI98-105P1RCP2DV1M

```
--D------ Majority
451 GLPSHF--HLQLSEIEFHEIIGSGSFGKVY
                                    CARP2/prot
58 ------LAKLETSNVIARGTYGTVY
                                    A.thaliana kinase2
83 -----LSQLFIGNKFASGAHSRIY
                                    A.thaliana3
Arabidopsis thaliana
496 ALPAPF - - HLCLAEIEFQESIGSGSFGKVY
                                    c.elegans kinase
81 - VRSEY - - EIDFNELEFGQTIGKGFFGEVK
                                    D.discoideum (A35670)
D.Discoideum (U01064)
496 GSDGAS - - PPASPSIIPRLRAIRLTPVDCG
                                    H.sapiens (Z48615)
496 GSDGAS--PPASPSIIPRLRAIRLTPVDCG
                                    Homo sapiens (Z48615)
335 QRDSSYYWEIEASEVMLSTRIGSGSFGTVY
                                    Human raf1(W13107)
335 QRDSSYYWEIEASEVMLSTRIGSGSFGTVY
                                    Human Raf1kinase(R98215)
Soybean kinase (M67449)
   KGIYRG-DVAVKILKRGDP--E----K-E Majority
            580
                                  600
479 KGRCRNKIVAIKR-YRAN----TYCSKSD
                                    CARP2/prot
77 KGIYDGQDVAVKVLDWEDDGNETTAKTATN
                                    A.thaliana kinase2
102 RGIYKQRAVAVKMVRIPTHKEETR--AKLE
                                    A.thaliana3
                                    Arabidopsis thaliana
c.elegans kinase
58 RGYWRETDVAIKIIYRDQ----FKTKSS
                                    D.discoideum (A35670)
157 LGIWNGYKVAIKILKNESISNDEK-----
                                    D.Discoideum (U01064)
524 GSSSGSSSGGSGTWSRGGPPKKEELVGGKK
                                    H.sapiens (Z48615)
524 GSSSGSSSGGSGTWSRGGPPKKEELVGGKK
                                    Homo sapiens (Z48615)
365 KGKWHG-DVAVKILKVVDPTPE-----Q
                                    Human raf1(W13107)
365 KGKWHG-DVAVKILKVVDPTPE----Q
                                    Human Raf1kinase(R98215)
172 HGVYKDEAVAVKIIMVPEDDGNGALASRLE
                                    Soybean kinase (M67449)
   KQ-FRNEVSVLSKLRHPNVVQFVGA-L---
                                    Majority
503 VDMFCREVSILCQLNHPCVIQFVGACL-N-
                                    CARP2/prot
107 RALFRQEVTVWHKLNHPNVTKFVGASMGTT
                                    A.thaliana kinase2
130 QQ-FKSEVALLSRLFHPNIVQFIAACK---
                                    A.thaliana3
Arabidopsis thaliana
548 TDMLCREVSILSRLAHPNVVAFVGTSL-D-
                                    c.elegans kinase
82 LVMFQNEVGILSKLRHPNVVQFLGACTAG-
                                    D.discoideum (A35670)
181 --- FIKEVSSLIKSHHPNVVTFMGA----
                                    D.Discoideum (U01064)
554 KGRTWGPSSTLQKERVGGEERLKG--LGEG
                                    H.sapiens (Z48615)
8554 KGRTWGPSSTLQKERVGGEERLKG--LGEG
                                    Homo sapiens (Z48615)
387 FQAFRNEVAVLRKTRHVNILLFMGY----
                                    Human raf1(W13107)
387 FQAFRNEVAVLRKTRHVNILLFMGY-----
                                    Human Raf1kinase(R98215)
202; KQ-FIREVTLLSRLHHQNVIKFSAACR---
                                    Soybean kinase (M67449)
```

Fig. 3G

Applicants: Jeyaseelan Raju Filed: Herewith Serial No.: N/A Title: Novel Cark Protein and Nucleic Acid Molecules and Uses

Therefor

Atty/Agent: Jean M. Silveri

Attorney Docket No.: MPI98-105P1RCP2DV1M

```
-----DPDHLCIVTEYLSGGSL Majority
                      650
                                660
137 NLNIRSADSKGSLPQQACCVVVEYLPGGTL A.thaliana kinase2
      -----KPPVYCIITEYMSQGNL A.thaliana3
240 ----- Arabidopsis thaliana
576 -------DPSQFAIITEFVENGSL c.elegans kinase
203 -----RIDPP--CIFTEYLQGGSL D.Discoideum (U01064)
582 SKQWSSSAPNLGKSPKHTPIAPGFASLNEM H.sapiens (Z48615)
582 SKQWSSSAPNLGKSPKHTPIAPGFASLNEM Homo sapiens (Z48615)
REHLH-ED-KFSLLP-----LKIALDI Majority
                     680
                                690
548 FSLLHEQK---RILD--LQSK--LIIAVDV CARP2/prot
167 KQHLI-RHKSKKLAFKAV----IKLALDL A.thaliana kinase2
173 RMYLN-KKEPYSLSIETV----LRLALDI A.thaliana3
240 ----- Arabidopsis thaliana
593 FRRENGERKNYRVMD - - PAFR - - LRISLDV c.elegans kinase
128 RQFLTDH---FNLLEQNPHIR--LKLALDI D.discoideum (A35670)
220 YDVLH - - IQKIKLNPLMM - - - - YKMIHDL D.Discoideum (U01064)
612 EEFAEAEDGGSSVPPSPYSTPSYLSVPLPA H.sapiens (Z48615)
612 EEFAEAEDGGSSVPPSPYSTPSYLSVPLPA Homo sapiens (Z48615)
430 YKHLHVQETKFQMFQ-----LIDIARQT Human raf1(W13107)
430 YKHLHVQETKFQMFQ-----LIDIARQT Human Raf1kinase(R98215)
245 RAYLH-KLEHQTISLQKL-----IAFALDI Soybean kinase(M67449)
  ARGMEYLH - - - AQPIIHRDLKSHNILLDE - Majority
           700
                     710
                                720
571 AKGMEYLHN - LTQPIIHRDLNSHNILLY - - CARP2/prot
191 ARGLSYLH - - - SEKIVHRDVKTENMLLDAQ A.thaliana kinase2
197 SRGMEYLH - - - SQGVIHRDLKSNNLLLNDE A.thaliana3
240 ----- Arabidopsis thaliana
619 ARGMRYLHESAAKPVIHRDLNSHNILIH - - c.elegans kinase
153 AKGMNYLHGW-TPPILHRDLSSRNILLDHN D.discoideum (A35670)
243 SLGMEHLH - - - SIQMLHRDLTSKNILLDEF D.Discoideum (U01064)
642 EPSPGARAPWEPTPSAPPARWGHG----AR H.sapiens (Z48615)
642 EPSPGARAPWEPTPSAPPARWGHG----AR Homo sapiens (Z48615)
453 AQGMDYLH - - - AKNIIHRDMKSNNIFLHEG Human raf1(W13107)
453 AQGMDYLH---AKNIIHRDMKSNNIFLHEG Human Raf1kinase(R98215)
269 ARGMEYIH - - - SQGVIHRDLKPENILINED Soybean kinase (M67449)
```

Fig. 3H

Applicants: Jeyaseelan Raju
Serial No.: N/A Filed: Herewith
Title: Novel Cark Protein and Nucleic Acid Molecules and Uses

Therefor

Atty/Agent: Jean M. Silveri

Attorney Docket No.: MPI98-105P1RCP2DV1M

```
598 - - - - - - - EDGHAVVADFGESRFLQS - CARP2/prot
 255 ----- W Arabidopsis thaliana
 647 -----ADGRSVVADFGESRFVCQ - c.elegans kinase
 182 IDPKNPLVSSRQDIKCKISDFGLSR-LKK- D.discoideum (A35670)
 668 RRCDLALLGC - - - - - ATLLGAVGLGAD - H. sapiens (Z48615)
 668 RRCDLALLGC-----ATLLGAVGLGAD- Homo sapiens (Z48615)
 480 LTVK------IGDFGLATVKSRW Human Raf1kinase(R98215)
 296 N H L K - - - - - - - - - I A D F G I A C - E E A - Soybean kinase (M67449)
   LADDQAADGTGTLRWMAPEVFI - - KGGPYS Majority
                        770
                                    780
 616 LDEDNMTKQPGNLRWMAPEVFT - - QCTRYT CARP2/prot
 233 LNPKDMTGRTGTLGYMAPEV-I--DGKPYN A.thaliana kinase2
 239 - QCREAKGNMGTYRWMAPEM-I--KEKPYT A.thaliana3
 267 - ADKQFAEEIQTRQYRAPEVILK - - - SGYS Arabidopsis thaliana
 665 REDENLTKQPGNLRWMAPEVFS - - QSGKYD c.elegans kinase
 210 EQASQMTQSVGCIPYMAPEVF - - - KGDSNS D.discoideum (A35670)
 283 LSDDMTLSGITNPRWRSPEL-T--KGLVYN D.Discoideum (U01064)
 690 VAEARAADGEEQRRWLDGLFFP -- RAGRFP H.sapiens (Z48615)
 690 VAEARAADGEEQRRWLDGLFFP -- RAGRFP Homo sapiens (Z48615)
 497 SGSQQVEQPTGSVLWMAPEVIRMQDNNPFS Human raf1(W13107)
 497 SGSQQVEQPTGSVLWMAPEVIRMQDNNPFS Human Raf1kinase(R98215)
 311 - SCDLLADDPGTYRWMAPEM-I--KRKSYG Soybean kinase(M67449)
   RKVDVYSFGLVLWELVTGELPFAHLNP-VQ Majority
             790
                                    810
 644 IKADVFSYALCLWEILTGEIPFAHLKP-AA CARP2/prot
 260 RRCDVYSFGICLWEIYCCDMPYPDLSF-VD A.thaliana kinase2
265 RKVDVYSFGIVLWELTTALLPFQGMTP-VQ A.thaliana3
 293 FSVDMWSFGCTAFELVTGDMLFAPKDGN - - Arabidopsis thaliana
 693 RKVDVFSFALVIWEIHTAELPFSHLKP-AA c.elegans kinase
 237 EKSDVYSYGMVL.FELLTSDEPQQDMKP-MK D.discoideum (A35670)
310 EKVDVYSFGLVVYEIYTGKIPFEGLDG-TA D.Discoideum (U01064)
718 RGLSPPARPHGRREDVGPGLGLAPSATLVS H.sapiens (Z48615)
718 RGLSPPARPHGRREDVGPGLGLAPSATLVS Homo sapiens (Z48615)
527 FQSDVYSYGIVLYELMTGELPYSHINNRDQ Human raf1(W13107)
527 FQSDVYSYGIVLYELMTGELPYSHINNRDQ Human Raf1kinase(R98215)
337 KKVDVYSFGLILWEMLTGTIPYEDMNP-IQ Soybean kinase(M67449)
```

Applicants: Jeyaseelan Raju Serial No.: N/A Filed: Herewith Title: Novel Cark Protein and Nucleic Acid Molecules and Uses

Therefor

Atty/Agent: Jean M. Silveri

Attorney Docket No.: MPI98-105P1RCP2DV1M

```
AAFAVAYGNARPPLPSDC - - - PAALSSLI Majority
             820
                        830
                                    840
673 AAADMAYHHIRPPIGYS----IPKPISSLL CARP2/prot
289 VSSAVVLHNLRPEIPRCC----PTALAGIM A.thaliana kinase2
294 AAFAVAEKNERPPLPASC----QPALAHLI A.thaliana3
321 ----- GYGEDEDHLA---------
                                      Arabidopsis thaliana
722 AAAEMTYKRGRPTLPNQPTAQFPAHILSLI c.elegans kinase
266 MAHLAAYESYRPPIPLTTSSKWKE----IL D.discoideum (A35670)
339 SAAKAAFENYRPAIPPDC----PVSLRKLI D.Discoideum (U01064)
748 LSSVSDCNSTRSLLRSDSDEAAPAAPSPPP H.sapiens (Z48615)
748 LSSVSDCNSTRSLLRSDSDEAAPAAPSPPP Homo sapiens (Z48615)
557 IIFMVGRGYASPDLSKLYKNC-PKAMKRLV Human raf1(W13107)
557 IIFMVGRGYASPDLSKLYKNC-PKAMKRLV Human Raf1kinase(R98215)
366 AAFAVVNKNSRPIIPSNC----PPAMRALI Soybean kinase(M67449)
   AQCWAPNPSKRPSFSEIV---LE---- Majority
             850
                                    870
699 IRGWNACPEGRPEFSEVVMK-LEECLCNIE CARP2/prot
315 KTCWDGNPQKRPEMKEVVKM-LEGV--DTS A.thaliana kinase2
320 KRCWSENPSKRPDFSNIVAV-LEKY--DEC A.thaliana3
331 ----- Arabidopsis thaliana
752 PQAWHPESSSRPDFVEIV------A c.elegans kinase
292 TQCWDSNPDSRPTFKQIIVH-LKE---- D.discoideum (A35670)
365 TKCWASDPSQRPSFTEILTE-LETM--KSK D.Discoideum (U01064)
778 SPP-APTPTPSPSTNPLVDLELESFKKDPR H.sapiens (Z48615)
778 SPP-APTPTPSPSTNPLVDLELESFKKDPR Homo sapiens (Z48615)
586 ADCVKKVKEERPLFPQIL------ Human raf1(W13107)
586 ADCVKKVKEERPLFPQIL------ Human Raf1kinase(R98215)
392 EQCWSLQPDKRPEFWQVVKI-LEQF--ES- Soybean kinase(M67449)
   ----P-SVTS--SLSL----TPS---- Majority
                        890
                                    900
728 L-MSPASSNSSGSLSP----SSSSDCLVNR CARP2/prot
342 K-GGGMIPE---------- A.thaliana kinase2
347 V-KEGLPLTSHASLTK----TKKA----- A.thaliana3
336 ----- Arabidopsis thaliana
771 L-LEPHVESTHTDISA----PSTV---- c.elegans kinase
315 -- MEDQGVSSFASVPV----QT----- D.discoideum (A35670)
392 F-IKQLSFLND--LIQ----NPD----- D.Discoideum (U01064)
807 QSLTPTHVTAACAVSRGHRRTPSDGALGQR H.sapiens (Z48615)
807 QSLTPTHVTAACAVSRGHRRTPSDGALGQR Homo sapiens (Z48615)
.604 -----RSIELLQHSLPK----INR Human Raf1kinase(R98215)
418 ----- SLASDGTLSL---- VPNP---- Soybean kinase(M67449)
```

Fig. 3J

Serial No.: N/A Filed: Herewith
Title: Novel Cark Protein and Nucleic Acid Molecules and Uses

Therefor

Atty/Agent: Jean M. Silveri

Attorney Docket No.: MPI98-105P1RCP2DV1M

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366	_	_	_	_	_	_	_	_	_	_			_	_	_	_	_	_	_	_	_	_	<u>-</u>	_	-	_	_	_	_	A.thaliana kinase2 A.thaliana3
349	_	_	-	K	S	K	D	Y	F	D :	R F	I G	D	L	K	R	Ι	R	R	L	_	_	_	_	ĸ.	v ·	W	Þ	т.	Arabidonsis thaliana
790	-	-	-	-	-	S	-	Q	L	T :	S () W	Ε	0	L	S	V	Α		P	P	P	A	S	K :	F	P	P	I	c.elegans kinase
331	_	-	-	-	-	-	-	-	-	_		-	_	_	_	-	_	_	_	_	_	_	_	_	_	_	_	_	_	D.discoideum (A35670)
408	-	-	-	_	_	-	-	-	-	_			<u>-</u>	-	-	<u>-</u>	_	-	-	-	-	-	-	-	-	-	-	-	-	D.Discoideum (U01064)
837	G	פ	D.	E	ב	A	G	H	G	P (G E	R	D	L	L	D	F	P	R	L	P	D	P	Q.	A I	L	F	P	A	H.sapiens (Z48615)
837 619	S	A	g	E	P	S	G T.	n H	B	P (A)	JI	אני יייו	עי	Д П	L	NI NI	r -	<u>ہ</u>	K -	_ L	₽	ַ	יי	Q.	A.	L,	F	P	A	Homo sapiens (Z48615) Human raf1(W13107)
619																														Human Raf1kinase(R98215)
432																														Soybean kinase(M67449)
																														·
,	_	_	_	-	_	_	_	-	_	-		-	L	T	_	D	P	_	K	G	P	<u>-</u>	_	-		_	-	_	D	Majority
									94	10								9	50									9	60	Majority
782	S	Q	S	A	G	Q	Y	S	S		- Ç	G	L	S	L	E	E	M	K	R	S	L	0 .	Y '	r i	Р	_	I	D	CARP2/prot A.thaliana kinase2
350	-	-	-	-	-	-	-	-	-			_	-	-	-	D	Q	S	R	G	C :	F	Ĉ.	-		_	_	_	_	A.thaliana kinase2
300	•••	_	-	_	_	-	-	-	-			_	***	Ι	L	ח	н	Τ.	K I	G	<i>C</i> '	v	ጥ	ς .	Т 9	₵.	_	-	g	A.thaliana?
372	D T.	K C	~ 7	- T	- U	~	- T	- 2	- 7		 m	-	L	L	I	D	K	Y	K	L	P:	E .	A]	E	A I	K I	E	F.	A	Arabidopsis thaliana c.elegans kinase
5.51	_	-	_	-	_	-	_	_	_			_	_	_	_	_	_	_	_	_	_						_	T	n	D.discoideum (A35670)
408	_	_	_	_	_	_	_	_	_			_	_	_	_	D	р [.]	- Y	- N 1	- N 1	N.	- · [. 1	N 1	- ·	ו ח	 R 1	- F: '	V	ח	D.Discoideum (U01064)
867	R	R	R	P :	Ρ	E.	F	P	\mathbf{G}	R 1	r	T	L	Т	F	A	P	R	P :	R	P :	A :	A :	S 1	R 1	P 1	R	т.	ח	H.sapiens (Z48615)
867	R	R	R	P	P	E	F.	P	G :	R I	? T	T	L	Т	F.	A	P	R	P	R :	P	A Z	A :	S I	R I	P 1	R :	L	D	Homo sapiens (Z48615)
636	~	-	-	-	-	-	-	-	-	- 1	7 C	T	L	T	T	S	P	-	-	-	-					- •	-	-	-	Human raf1(W13107)
636 432	_	_	<u>-</u>	<u>-</u>	-	_	_	_	_	- 1	r C	Т	L	T	T	S	P	 12 ·	- ·	-		- :			- :		-		- -	Human Raf1kinase(R98215)
434	_	_	_	_	_	_	_	_	_			_	-	C	₩.	ע.	n.	Κ.	K (. خی	ا ما	ا يا	H 1	Ν.	L (ו ג	K .	L	Ġ	Soybean kinase(M67449)
	P	F	_	-	_	-	_	_	_	- 1	<u> </u>	Ι	-	P	-	-	-]	P	_	_						. .		_	-	Majority
									97	0								9	80									9	- 90	
																														CARP2/prot
380	P:	F	-	_	-	-		-	- :	5 5	S	S	V	P	V	N					- -							_	_	A thaliana?
391	E :	F	-	-	-	- :	L '	T :	P :	ΙΙ	Ε	F	A	P	E]	K :	R I	Ρ'	T A	A (Q (2 (3 -					- :	L	Arabidopsis thaliana c.elegans kinase
333	Τ. Τ.	C.	_	<u> </u>	_	<u> </u>	<u> </u>		- (- 1	j) T	. V	T	N	-	_	-		-										-		c.elegans kinase
422		G	_					_ `		v ,	. –	_	_	_	_	_	•	-			- '			•	-					D.discoideum (A35670) D.Discoideum (U01064)
		W :	K :	<u>ر</u> ا	V	S I	F (G 1	R '	r I	T	I	S	P	P	S 1	R 1	P 1	י ם	r 1	PI	3 5	3 I	· (; ;) I	Р 9	s 1	J	H.sapiens (Z48615)
897	P	W.	K :	L	V	S	F (G 1	R :	r I	T	Ι	S	P	P	S	R I	P 1	D :	r i	PI	3	5 1	9	;)]	P (s 1	V	Homo sapiens (Z48615)
644	-	- :	R :	L]	P 1	V I	F																							Human raf1(W13107)
644									_			_	_	_						_										Human Raf1kinase(R98215)
448	צ.	L	-	- '	-	- ·		- '	- }	iς	N	S	G	₽ `	v :	נ פ	K I	?]	K I					-	-				-	Soybean kinase(M67449)

Fig. 3K

Applicants: Jeyaseelan Raju Serial No.: N/A Filed: Herewith Title: Novel Cark Protein and Nucleic Acid Molecules and Uses

Therefor

Atty/Agent: Jean M. Silveri

Attorney Docket No.: MPI98-105P1RCP2DV1M

	-	-	-	_	-	_	-	_	_	_	_	-	-	_	_	_	_	-	_	-	-	_	-		-	-	_	Majority
									1	0,0	0								1	01	.0							
	E	-	-	-	-	-	-	-	-	-	-	-	-	-	_	-	_	_	_	-	_	-	-	-	D	S	S	CARP2/prot
364 390	_	_	_	_	-	_	_	_	_	_	_	_	_	_	_	_	_	_	_	_	_	_	_	_	_	_	A	A.thaliana kinase2 A.thaliana3
412	D	H	P	W	M	N	٧	T	T	Q	N	D	A	E	N	V	D	D	Q	M	N	N	L	H	I	K	G	Arabidopsis thaliana
841																												c.elegans kinase
337	_	-	-	-	-	-	-	-	-	-	-	-	-	_	-	_	-	_	-	-	-	_	-	_	-	-	A	D.discoideum (A35670)
422																												D.Discoideum (U01064)
927	Q	P	T	L	L	D	M	D	M	E	G	Q	N	Q	D	S	T	V	P	L	C	G	A	H	G	S	H	H.sapiens (Z48615)
927	Q	P	T	L	L	D	M	D	M	E	G	Q	N	Q	D	S	T	V	P	L	C	G	A	Н	G	S	H	Homo sapiens (Z48615)
648												_		~														Human raf1(W13107)
648																												<pre>Human Raf1kinase(R98215)</pre>
462	_	_	_	_	-	_	_	-	-	_	-	-	_	_	_	_	_	_	_	-	_	_	_	_	_	_	T	Soybean kinase (M67449)

Fig. 3L

```
Atty/Agent: Jean M. Silveri
                Attorney Docket No.: MPI98-105P1RCP2DV1M
                                  16/35
Protein Family / Domain Matches, HMMer version 2
Searching for complete domains
hmmpfam - search a single seq against HMM database
HMMER 2.1.1 (Dec 1998)
Copyright (C) 1992-1998 Washington University School of Medicine
HMMER is freely distributed under the GNU General Public License (GPL).
                             ------------
HMM file:
                           /prod/ddm/seqanal/PFAM/pfam3.4/Pfam
Sequence file:
                           /tmp/orfanal.13255.aa
_ _ _ _ _ _ _ _
 Query: sequence13252
Scores for sequence family classification (score includes all domains):
Model
         Description
                                                          Score
                                                                   E-value N
          PF00023 Ank repeat
ank
                                                          207.5
                                                                     2e-58
pkinase PF00069 Eukaryotic protein kinase domain
                                                          201.9
                                                                   9.9e-57
Parsed for domains:
Model
         Domain seq-f seq-t
                                 hmm-f hmm-t
                                                   score E-value
            1/9
                           99 ..
                                                               12
ank
                     66
                                           33 []
                                                     3.2
ank
            2/9
                    100
                          132 ..
                                          33 []
                                                    34.0
                                                          3.4e-06
                          165 ..
ank
            3/9
                    133
                                          33 []
                                                    44.5
                                                          2.4e-09
ank
            4/9
                    168
                          198 ..
                                          33 []
                                                    34.6
                                                          2.3e-06
            5/9
ank
                    199
                          233 ..
                                     1
                                          33 []
                                                    28.1
                                                          0.00021
            6/9
 ank
                    234
                          268 ...
                                     1
                                          33 []
                                                    15.4
                                                             0.28
                          302 ..
            7/9
                                           33 []
 ank
                    269
                                                    20.6
                                                            0.037
                          338 ..
 ank
            8/9
                    306
                                           33 []
                                                     3.3
                                           33 []
                          371 ...
                                     1
                                                          1.3e-06
 ank
            9/9
                    339
                                                    35.4
            1/1
                    463
                          716 ...
                                     1
                                          273 [.
                                                   201.9 9.9e-57
 pkinase
 Alignments of top-scoring domains:
 ank: domain 1 of 9, from 66 to 99: score 3.2, E = 12
                    *->nGnTPLHlAaryg.nvevvklLLehGAdvnartk<-*
                       nG++ LH1+++ g++ + ++ L+ +G + t+
   sequence13
                 66
                       NGLSLLHLCCICGGKKSHIRTLMLKGLRPSRLTR
 ank: domain 2 of 9, from 100 to 132: score 34.0, E = 3.4e-06
                    *->nGnTPLHlAarygnvevvklLLehGAdvnartk<-*
                       nG+T+LHlA++++n e++ LL GAd+ + +
   sequence13
                100
                       NGFTALHLAVYKDNAELITSLLHSGADIQQVGY
                                                             132
 ank: domain 3 of 9, from 133 to 165: score 44.5, E = 2.4e-09
                    *->nGnTPLHlAarygnvevvklLLehGAdvnartk<-*
                        G+T+LH+A ++g++e +++LL+hGA+vn+++
   sequence13
                133
                       GGLTALHIATIAGHLEAADVLLQHGANVNIQDA
                                                             165
 ank: domain 4 of 9, from 168 to 198: score 34.6, E = 2.3e-06
                    *->nGnTPLHlAarygnvevvklLLehGAdvnartk<-*
                         +TPLH+Aa+yg+ +v +lLL+ GAdvn+
   sequence13
                168
                       --FTPLHIAAYYGHEQVTRLLLKFGADVNVSGE
                                                              198
 ank: domain 5 of 9, from 199 to 233: score 28.1, E = 0.00021
                    *->nGnTPLHlAarygnvevvklLLe..hGAdvnartk<-*
                        G+ PLH1A+ +g ++++klL+e++ Advna+++
   sequence13
                199
                       VGDRPLHLASAKGFLNIAKLLMEegSKADVNAQDN
                                                               233
```

Filed: Herewith

Title: Novel Cark Protein and Nucleic Acid Molecules and Uses

Applicants: Jeyaseelan Raju

Serial No.: N/A

Fig. 4A

ank: domain 6 of 9, from 234 to 268: score 15.4, E = 0.28 *->nGnTPLH1AarygnvevvklLLe..hGAdvnartk<-* + PLH ++r g+ ++vk+LL+++ sequence13 234 EDHVPLHFCSRFGHHDIVKYLLQsdLEVQPHVVNI 268 ank: domain 7 of 9, from 269 to 302: score 20.6, E = 0.037 *->nGnTPLHlAarygnvevvklLLe.hGAdvnartk<-* G+TPLH1A+++g+ ev+k ++ G+ sequence13 269 YGDTPLHLACYNGKFEVAKEIIQiSGTESLTKEN 302 ank: domain 8 of 9, from 306 to 338: score 3.3, E = 11 *->nGnTPLHlAaryg.nvevvklLLe.hGAdvnartk<-** T+ H A+ yg+++++vk+LL+++ ++n + + --ETAFHSACTYGKSIDLVKFLLDqNVININHQGR sequence13 306 338 ank: domain 9 of 9, from 339 to 371: score 35.4, E = 1.3e-06 *->nGnTPLHlAarygnvevvklLLehGAdvnartk<-* +G+T LH A+++g++++v++LL++GAd n + sequence13 339 DGHTGLHSACYHGHIRLVQFLLDNGADMNLVAC 371 pkinase: domain 1 of 1, from 463 to 716: score 201.9, E = 9.9e-57 *->yelleklGeGsfGkVykakhktgkivAvKilk.....kesls..lr +e++e++G+GsfGkVyk++ + +kivA+K + ++ +k++++ sequence13 463 IEFHEIIGSGSFGKVYKGRCR-NKIVAIKRYRantycsKSDVDmfCR 508 EiqilkrlsHpNIvrllgvfedtddhlylvmEymegGdLfdylrrng.pl E++il +l+Hp ++++ g++ ++++ + +v++y gG+Lf++l+++++ 1 sequence13 509 EVSILCQLNHPCVIQFVGAVLNDPSQFAIVTQYISGGSLFSLLHEQKrIL 558 sekeakkialQilrGleYLHsng..ivHRDLKpeNILldengtvKiaDFG + + + ia ++++G+eYLH+ ++i+HRDL + NILl e+g+ +aDFG sequence13 559 DLQSKLIIAVDVAKGMEYLHNLTqpIIHRDLNSHNILLYEDGHAVVADFG 608 LArll....eklttfvGTpwYmmAPEvileg.rgysskvDvWSlGviLy +r+1++ ++ ++t G +++m APEv + + y+ k+Dv S+ sequence13 609 ESRFLqsldeDNMTKQPGNLRWM-APEV-FTQcTRYTIKADVFSYALCLW 656 ElltggplfpgadlpaftggdevdqliifvlklPfsdelpktridpleel E+ltq ++Pf + 657 EILTG-----LKPAAAA 674 sequence13 frikkr..rlplpsncSeelkdLlkkcLnkDPskRpGsatakeil<-* ++ +++ r+p+ ++++++ +Ll + +n P+ Rp sequence13 675 ADMAYHhiRPPIGYSIPKPISSLLIRGWNACPEGRP---EFSEVV 716

Filed: Herewith

Title: Novel Cark Protein and Nucleic Acid Molecules and Uses

17/35

Attorney Docket No.: MPI98-105P1RCP2DV1M

Applicants: Jeyaseelan Raju

Atty/Agent: Jean M. Silveri

Serial No.: N/A

Fig. 4B

Filed: Herewith Title: Novel Cark Protein and Nucleic Acid Molecules and Uses

Therefor

Atty/Agent: Jean M. Silveri

Attorney Docket No.: MPI98-105P1RCP2DV1M

gtc	gacc	cac (gcgt	ccgg	tg a	aggg	cagc	a gc	acag	gaga	aaa	gcaa	aga	cttc	tttaaa	60
atg Met 1	ggg Gly	aat Asn	tac Tyr	aaa Lys 5	tcc Ser	aga Arg	cca Pro	aca Thr	cag Gln 10	act Thr	tgt Cys	tct Ser	gat Asp	gaa Glu 15	tgg Trp	108
aag Lys	aag Lys	aaa Lys	gtt Val 20	agt Ser	gaa Glu	tct Ser	tac Tyr	gct Ala 25	att Ile	atc Ile	ata Ile	gaa Glu	agg Arg 30	ctg Leu	gag Glu	156
gat Asp	aac Asn	ctg Leu 35	cag Gln	atc Ile	aaa Lys	gaa Glu	aat Asn 40	gaa Glu	ttt Phe	caa Gln	gaa Glu	cta Leu 45	agg Arg	cac His	atc Ile	204
ttt Phe	ggc Gly 50	tct Ser	gat Asp	gaa Glu	gcc Ala	ttc Phe 55	agt Ser	gaa Glu	gtc Val	agt Ser	tta Leu 60	aat Asn	tac Tyr	cgc Arg	aca Thr	252
gag Glu 65	cgt Arg	ggc Gly	ctg Leu	tcc Ser	ctg Leu 70	cta Leu	cac His	ctc Leu	tgc Cys	tgt Cys 75	gtc Val	tgt Cys	ggc Gly	ggc Gly	aac Asn 80	300
aag Lys	tca Ser	cat His	atc Ile	cgt Arg 85	gcc Ala	ctt Leu	atg Met	tta Leu	aaa Lys 90	Gly	ctc Leu	cgt Arg	cca Pro	tcc Ser 95	aga Arg	348
ctg Leu	acg Thr	aga Arg	aat Asn 100	ggg Gly	ttt Phe	cca Pro	gct Ala	ctg Leu 105	cac His	ctg Leu	gcc Ala	gtt Val	tac Tyr 110	aag Lys	gac Asp	396
agc Ser	ccg Pro	gaa Glu 115	ctt Leu	atc Ile	act Thr	tca Ser	ctg Leu 120	ttg Leu	cac His	agc Ser	gga Gly	gca Ala 125	gat Asp	gtt Val	cag Gln	444
caa Gln	gtg Val 130	gga Gly	tac Tyr	ggt Gly	ggc Gly	ctc Leu 135	aca Thr	gcc Ala	ctc Leu	cac His	ata Ile 140	gct Ala	gca Ala	ata Ile	gct Ala	492
gga Gly 145	cac His	cca Pro	gag Glu	gct Ala	gca Ala 150	gaa Glu	gtg Val	ctg Leu	cta Leu	caa Gln 155	cat His	ggg Gly	gcc Ala	aat Asn	gtg Val 160	540
aat Asn	vaı	caa Gln	gat Asp	gcc Ala 165	gtc Val	ttc Phe	ttc Phe	acc Thr	cca Pro 170	ctg Leu	cac His	att Ile	gca Ala	gcc Ala 175	tac Tyr	588
tat Tyr	ggg	cac His	gag Glu 180	cag Gln	gta Val	acc Thr	agt Ser	gtc Val 185	ctt Leu	ttg Leu	aag Lys	ttt Phe	ggt Gly 190	gct Ala	gat Aspp	636
gtc Val	aat Asn	gta Val 195	agc Ser	ggt Gly	gaa Glu	gtt Val	ggg Gly 200	gac Asp	agg Arg	cct Pro	ctg Leu	cac His 205	ctg Leu	gcc Ala	tct Ser	684

Applicants: Jeyaseelan Raju Serial No.: N/A Filed: Herewith

Title: Novel Cark Protein and Nucleic Acid Molecules and Uses

Therefor

Atty/Agent: Jean M. Silveri

Attorney Docket No.: MPI98-105P1RCP2DV1M

19/35

								9/35								
gca Ala	aag Lys 210	ggc Gly	ttc Phe	ttc Phe	aac Asn	att Ile 215	gtg Val	aaa Lys	ctc Leu	ctg Leu	gta Val 220	gaa Glu	gaa Glu	ggg Gly	agc Ser	732
aaa Lys 225	gca Ala	gat Asp	gtg Val	aac Asn	gct Ala 230	cag Gln	gac Asp	aat Asn	gaa Glu	gac Asp 235	cac His	gtc Val	cct Pro	ctg Leu	cac His 240	780
ttc Phe	tgt Cys	tct Ser	cga Arg	ttt Phe 245	gga Gly	cac His	cac His	aat Asn	ata Ile 250	gtg Val	agc Ser	tac Tyr	ctg Leu	ctc Leu 255	cag Gln	828
agt Ser	gac Asp	tta Leu	gag Glu 260	gtc Val	cag Gln	cct Pro	cac His	gtc Val 265	att Ile	aac Asn	atc Ile	tat Tyr	ggt Gly 270	gac Asp	act Thr	876
cct Pro	ttg Leu	cac His 275	ctg Leu	gca Ala	tgc Cys	tac Tyr	aat Asn 280	gga Gly	aat Asn	ttt Phe	gaa Glu	gtt Val 285	gcc Ala	aag Lys	gaa Glu	924
att Ile	gtc Val 290	cag Gln	gta Val	aca Thr	gga Gly	act Thr 295	gaa Glu	agt Ser	ctg Leu	act Thr	aag Lys 300	gaa Glu	aac Asn	atc Ile	ttc Phe	972
agc Ser 305	gag Glu	aca Thr	gct Ala	ttt Phe	cac His 310	agt Ser	gct Ala	tgt Cys	acc Thr	tat Tyr 315	ggc Gly	aag Lys	aac Asn	att Ile	gac Asp 320	1020
ctg Leu	gtc Val	aaa Lys	ttt Phe	ctt Leu 325	ctt Leu	gat Asp	cag Gln	aat Asn	gct Ala 330	gtg Val	aac Asn	att Ile	aac Asn	cac His 335	cga Arg	1068
gga Gly	aga Arg	gat Asp	ggg Gly 340	cac His	aca Thr	gga Gly	ttg Leu	cac His 345	tct Ser	gct Ala	tgc Cys	tac Tyr	cac His 350	ggc Gly	cat His	1116
atc Ile	cgc Arg	ctg Leu 355	gtt Val	cag Gln	ttc Phe	cta Leu	ctt Leu 360	gat Asp	aat Asn	ggt Gly	gca Ala	gat Asp 365	atg Met	aat Asn	ctt Leu	1164
gtc Val	gct Ala 370	tgt Cys	gat Asp	ccc Pro	agc Ser	agg Arg 375	tct Ser	agt Ser	ggt Gly	gaa Glu	aaa Lys 380	gat Asp	gag Glu	cag Gln	aca Thr	1212
tgt Cys 385	ttg Leu	atg Met	tgg Trp	gct Ala	tac Tyr 390	gag Glu	aaa Lys	gga Gly	cat His	gat Asp 395	gcc Ala	att Ile	gtt Val	aca Thr	ctc Leu 400	1260
ctg Leu	aag Lys	cac His	tac Tyr	aag Lys 405	aga Arg	ccc Pro	cag Gln	gag Glu	gag Glu 410	ctg Leu	cca Pro	tgt Cys	aac Asn	gaa Glu 415	tat Tyr	1308
tcc Ser	cag Gln	cct Pro	gga Gly 420	gga Gly	gat Asp	ggc Gly	Ser	tat Tyr 425	Val	tct Ser	gtt Val	cct Pro	tcc Ser 430	ccc Pro	ttg Leu	1356

Fig. 5B

Serial No.: N/A Filed: Herewith
Title: Novel Cark Protein and Nucleic Acid Molecules and Uses

Therefor

Atty/Agent: Jean M. Silveri

Attorney Docket No.: MPI98-105P1RCP2DV1M

							21	JISS								
										gca Ala						1404
										caa Gln						1452
										ggg Gly 475						1500
										cga Arg						1548
										tgc Cys						1596
										cag Gln						1644
ctg Leu	gat Asp 530	gac Asp	ccc Pro	agt Ser	cag Gln	ttt Phe 535	gcc Ala	att Ile	gtc Val	act Thr	cag Gln 540	tac Tyr	att Ile	tca Ser	gga Gly	1692
										aag Lys 555						1740
										gcc Ala						1788
										cgc Arg						1836
										gtg Val						1884
										gac Asp						1932
										gtg Val 635						1980
										tac Tyr						2028

Applicants: Jeyaseelan Raju Serial No.: N/A Filed: Herewith

Title: Novel Cark Protein and Nucleic Acid Molecules and Uses

Therefor

Atty/Agent: Jean M. Silveri

Attorney Docket No.: MPI98-105P1RCP2DV1M

21/35

							2	1/35								
gag Glu	ctc Leu	ctc Leu	act Thr 660	gga Gly	gaa Glu	att Ile	cca Pro	ttc Phe 665	gct Ala	cat His	ctc Leu	aag Lys	cca Pro 670	gcc Ala	gct Ala	2076
gca Ala	gca Ala	gca Ala 675	gat Asp	atg Met	gcg Ala	tat Tyr	cac His 680	cac His	atc Ile	aga Arg	ccg Pro	ccc Pro 685	atc Ile	ggc Gly	tat Tyr	2124
tcc Ser	atc Ile 690	ccc Pro	aag Lys	ccc Pro	atc Ile	tca Ser 695	tcc Ser	ctg Leu	ctg Leu	ata Ile	cgg Arg 700	ggc Gly	tgg Trp	aat Asn	gca Ala	2172
tgt Cys 705	cct Pro	gaa Glu	gga Gly	cga Arg	cca Pro 710	gag Glu	ttc Phe	tct Ser	gaa Glu	gtc Val 715	gtt Val	agc Ser	aaa Lys	ctg Leu	gag Glu 720	2220
gag Glu	tgc Cys	cta Leu	tgc Cys	aat Asn 725	gtg Val	gag Glu	ctc Leu	atg Met	tct Ser 730	cca Pro	gca Ala	tca Ser	agt Ser	aac Asn 735	agc Ser	2268
agt Ser	ggc Gly	tct Ser	ctg Leu 740	tca Ser	cct Pro	tcc Ser	tct Ser	tct Ser 745	tcc Ser	gat Asp	tgc Cys	ctg Leu	ctg Leu 750	agc Ser	cgg Arg	2316
gga Gly	ggg Gly	cct Pro 755	ggc Gly	cgg Arg	agc Ser	cac His	gtg Val 760	gca Ala	gcc Ala	tta Leu	cgg Arg	agc Ser 765	cgt Arg	ttt Phe	gag Glu	2364
ttg Leu	gag Glu 770	tat Tyr	gcc Ala	cta Leu	aat Asn	gca Ala 775	agg Arg	tcc Ser	tat Tyr	gct Ala	ggg Gly 780	tgg Trp	tcc Ser	caa Gln	agt Ser	2412
gtt Val 785	gga Gly	aca Thr	cac His	tct Ser	aat Asn 790	ccg Pro	ggc Gly	ctg Leu	tct Ser	ttg Leu 795	gag Glu	gag Glu	atg Met	aat Asn	agg Arg 800	2460
agc Ser	acc Thr	cag Gln	tat Tyr	tca Ser 805	act Thr	gtt Val	gac Asp	aaa Lys	tac Tyr 810	ggc Gly	tat Tyr	gtg Val	tct Ser	gat Asp 815	ccc Pro	2508
atg Met	agc Ser	ctg Leu	acg Thr 820	cac His	ctt Leu	cac His	tcc Ser	cgc Arg 825	caa Gln	gac Asp	gac Asp	agc Ser	aac Asn 830	ttt Phe	gag Glu	2556
gac Asp	agc Ser	aac Asn 835	tgac	aggt	ct g	gcat	acad	cc ta	aggg	gcgt	cto	ccca	itca			2605
ggct	gaca	gc a	ataa	tttt	a co	cato	racac	r act	tact	tee	aatt	2+22		act o	ccctc	2665
tgao	gttt	ct t	caaa	tcat	c th	actt	atto	taa	acto	att	taat	tooo	itt c	rtage	ggaca	2003 2725
ggct	ttga	ct c	atgo	caag	ıc ct	gaaq	rtgto	aaa	gaac	aga	taca	gaat	at o	rcato	gggaa	2785
ttgt	tctt	ag t	ttga	tatt	t aa	agco	ctta	att	gcct	ggg	gcto	ggat	tc a	aato	tgtgt	2845
agat	agct	gg g	ttga	ccct	t at	gtat	ttgt	aga	ccaa	act	gtgt	gggc	tt c	tgtt	tgagg	2905
gtct	cctg	rtt g	ggtt	tctt	a aa	aaca	agct	ggc	tgat	tta	tctc	ctgt	tg c	cttt	gttgt	2965
tact	tctg	rtg a	ttaa	agto	t ct	tcgg	rtgat	: cta	gaaa	aaa	aaaa	aaaa	aa a	ıgggc	ggccg	3025
С									~ [- D						3026

Fig. 5D

```
Atty/Agent: Jean M. Silveri
                  Attorney Docket No.: MPI98-105P1RCP2DV1M
                                      22/35
Searching for complete domains in PFAM
hmmpfam - search a single seq against HMM database
HMMER 2.1.1 (Dec 1998)
Copyright (C) 1992-1998 Washington University School of Medicine
HMMER is freely distributed under the GNU General Public License (GPL).
HMM file:
                           /prod/ddm/seqanal/PFAM/pfam4.4/Pfam
/prod/ddm/wspace/orfanal/oa-script.11086.seq
Sequence file:
- - - - - - - - -
 Query: ratCARKpro
Scores for sequence family classification (score includes all domains):
Model Description
                                                                Score
                                                                          E-value N
ank
           Ank repeat
                                                                 212.7
                                                                           5.5e-60 9
pkinase
           Eukaryotic protein kinase domain
                                                                 206.4
                                                                           4.3e-58 1
Parsed for domains:
Model Domain seq-f seq-t hmm-f hmm-t
                                                       score E-value
           1/9
                                                                48
ank
                     66
                            99 ..
                                    1 33 []
                                                        1.2

      132
      1
      33 []

      165
      1
      33 []

      198
      1
      33 []

      233
      1
      33 []

      264
      1
      33 []

      302
      1
      33 []

      338
      1
      33 []

      371
      1
      33 []

      716
      1
      273. [.

ank
           2/9
                     100
                                                        28.6 0.00014
ank
           3/9
                     133
                                                        49.2
                                                               9.1e-11
                                                        31.9 1.4e-05
ank
           4/9
                     168
ank
           5/9
                     199
                                                        28.4 0.00017
ank
            6/9
                     234
                                                        12.6
                                                                    2.4
ank
            7/9
                     269
                                                        23.1
                                                                 0.0064
ank
            8/9
                     306
                                                        11.2
                                                                   3.5
ank
            9/9
                     339
                                                         36.4 6.5e-07
pkinase
            1/1
                     463
                            716 ..
                                        1
                                             273.[.
                                                       206.4 4.3e-58
Alignments of top-scoring domains:
ank: domain 1 of 9, from 66 to 99: score 1.2, E = 48
                      *->nGnTPLHlAaryg.nvevvklLLehGAdvnartk<-*
                         +G++ LH1++ g+n + +++L+ +G + t+
  ratCARKpro
                  66
                        RGLSLLHLCCVCGgNKSHIRALMLKGLRPSRLTR
                                                                    99
ank: domain 2 of 9, from 100 to 132: score 28.6, E = 0.00014
                      *->nGnTPLHlAarygnvevvklLLehGAdvnartk<-*
                         nG+ +LH1A+++++ e++ LL GAdv + +
  ratCARKpro
                        NGFPALHLAVYKDSPELITSLLHSGADVQQVGY
                 100
ank: domain 3 of 9, from 133 to 165: score 49.2, E = 9.1e-11
                     *->nGnTPLHlAarygnvevvklLLehGAdvnartk<-*
                          G+T+LH+Aa++g+e++LL+hGA+vn+++
  ratCARKpro
                 133
                        GGLTALHIAAIAGHPEAAEVLLQHGANVNVQDA
                                                                   165
ank: domain 4 of 9, from 168 to 198: score 31.9, E = 1.4e-05
                     *->nGnTPLHlAarygnvevvklLLehGAdvnartk<-*
                           +TPLH+Aa+yg+ +v +LL+ GAdvn+ +
  ratCARKpro
                 168
                        --FTPLHIAAYYGHEQVTSVLLKFGADVNVSGE
ank: domain 5 of 9, from 199 to 233: score 28.4, E = 0.00017
                     *->nGnTPLHlAarygnvevvklLLe..hGAdvnartk<-*
                         G+ PLH1A+ +g ++vk1L+e++ Advna+++
  ratCARKpro
                199
                        VGDRPLHLASAKGFFNIVKLLVEegSKADVNAQDN
                                                                     233
```

Filed: Herewith

Title: Novel Cark Protein and Nucleic Acid Molecules and Uses

Applicants: Jeyaseelan Raju

Serial No.: N/A

Fig. 6A

```
23/35
ank: domain 6 of 9, from 234 to 264: score 12.6, E = 2.4
                   *->nGnTPLH1Aarygnvevvk1LLehGAdvnartk<-*
                        + PLH ++r q+ ++v +LL+ +d ++
 ratCARKpro
                     EDHVPLHFCSRFGHHNIVSYLLQ--SDLEVQPH
               234
ank: domain 7 of 9, from 269 to 302: score 23.1, E = 0.0064
                   *->nGnTPLHlAarygnvevvklLLe.hGAdvnartk<-*
                       G+TPLHlA+++gn ev+k ++ G+
 ratCARKpro
               269
                     YGDTPLHLACYNGNFEVAKEIVQvTGTESLTKEN
                                                           302
ank: domain 8 of 9, from 306 to 338: score 11.2, E = 3.5
                   *->nGnTPLHlAatyg.nvevvklLLe.hGAdvnartk<-*
                         T+ H A+ yg+n+++vk+LL+++ ++n r +
  ratCARKpro
               306
                     --ETAFHSACTYGKNIDLVKFLLDqNAVNINHRGR
                                                             338
ank: domain 9 of 9, from 339 to 371: score 36.4, E = 6.5e-07
                   *->nGnTPLHlAarygnvevvklLLehGAdvnartk<-*
                      +G+T LH A+++g++++v++LL++GAd n +
  ratCARKpro
               339
                      DGHTGLHSACYHGHIRLVQFLLDNGADMNLVAC
                                                           371
pkinase: domain 1 of 1, from 463 to 716: score 206.4, E = 4.3e-58
                   *->yelleklGeGsfGkVykakhktgkivAvKilk.....kesls..lr
                      +e++e++G+GsfGkVyk++ + +kivA+K + ++ +k++++
  ratCARKpro
               463
                      IEFHEIIGSGSFGKVYKGRCR-NKIVAIKRYRantycsKSDVDmfCR 508
                   EiqilkrlsHpNIvrllgvfedtddhlylvmEymegGdLfdylrrng.pl
                   E++il +l+Hp +v++ g++ d+++ + +v++y gG+lf++l+++++ l
  ratCARKpro
               509 EVSILCQLNHPCVVQFVGACLDDPSQFAIVTQYISGGSLFSLLHEQHrIL 558
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                   + + + ia ++++G+eYLHs ++i+HRDL + NILl e+g+ +aDFG
  ratCARKpro
               559 DLQSKLIIAVDVAKGMEYLHSLTQDIIHRDLNSHNILLYEDGHAVVADFG 608
                   LArll.....eklttfvGTpwYmmAPEvileg.rgysskvDvWSlGviLy
                    +r+l++ ++ ++t
                                    G +++m APEv
                                                + + y+ k+Dv S+
               609 ESRFLqsldeDNMTKQPGNLRWM-APEV-FTQcTRYTIKADVFSYSLCLW 656
  ratCARKpro
                   ElltggplfpgadlpaftggdevdgliifvlklPfsdelpktridpleel
                   Elltg
                                                  ++Pf +
  ratCARKpro
              -657 ELLTG----
                                -----LKPAAAA 674
                   frikkr..rlplpsncSeelkdLlkkcLnkDPskRpGsatakeil<-*
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  ratCARKpro
              675 ADMAYHhirPPIGYSIPKPISSLLIRGWNACPEGRP---EFSEVV
                                                                    716
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Filed: Herewith

Title: Novel Cark Protein and Nucleic Acid Molecules and Uses

Attorney Docket No.: MPI98-105P1RCP2DV1M

Applicants: Jeyaseelan Raju

Atty/Agent: Jean M. Silveri

Serial No.: N/A

Fig. 6B

Attorney Docket No.: MPI98-105P1RCP2DV1M 24/35 GAP of: AAAa006_n check: 4956 from: 1 to: 3025 cark (analysis only) - Import - complete to: BAAa006_n check: 6389 from: 1 to: 3026 Rat CARK cDNA (analysis only) - Import - complete Symbol comparison table: /ddm local/gcg/gcg 9.1/gcgcore/data/rundata/ nwsgapdna.cmp CompCheck: 8760 Gap Weight: 12 Average Match: 10.000 Length Weight: 4 Average Mismatch: 0.000 Quality: 24376 Length: 3045 Ratio: 8.058 Gaps: 14 Percent Similarity: 82.169 Percent Identity: 82.169 Match display thresholds for the alignment(s): = IDENTITY 5 1 AAAa006_n x BAAa006_n 1 gtcgacccacgcgtccg.. ..gccctggagaaaggaagaaa 37 1 GTCGACCCACGCGTCCGGTGAAGGCCAGCAGCACAGGAGAAAAGCAAAGA 50 38 cttataataaatgggaaattataaatctagaccaacccaaacttgtactg 87 51 CTTCTTTAAAATGGGGAATTACAAATCCAGACCAACACAGACTTGTTCTG 100 88 atgaatggaagaaaaagtcagtgaatcatatgttatcacaatagaaaga 137 101 ATGAATGGAAGAAGAAGTTAGTGAATCTTACGCTATTATCATAGAAAGG 150 138 ttagaagatgacctgcagatcaaggaaaaagaactgacagaactaaggaa 187 151 CTGGAGGATAACCTGCAGATCAAAGAAAATGAATTTCAAGAACTAAGGCA 200 188 tatatttggctctgatgaagccttcagtaaagtcaatttaaattaccgca 237 201 CATCTTTGGCTCTGATGAAGCCTTCAGTGAAGTCAGTTTAAATTACCGCA 250 238 ctgaaaatgggctgtctctacttcatttatgttgcatttgtggaggcaag 287 251 CAGAGCGTGGCCTGTCCCTGCTACACCTCTGCTGTGTCTGTGGCGGCAAC 300

Filed: Herewith

Title: Novel Cark Protein and Nucleic Acid Molecules and Uses

Applicants: Jeyaseelan Raju

Atty/Agent: Jean M. Silveri

Serial No.: N/A

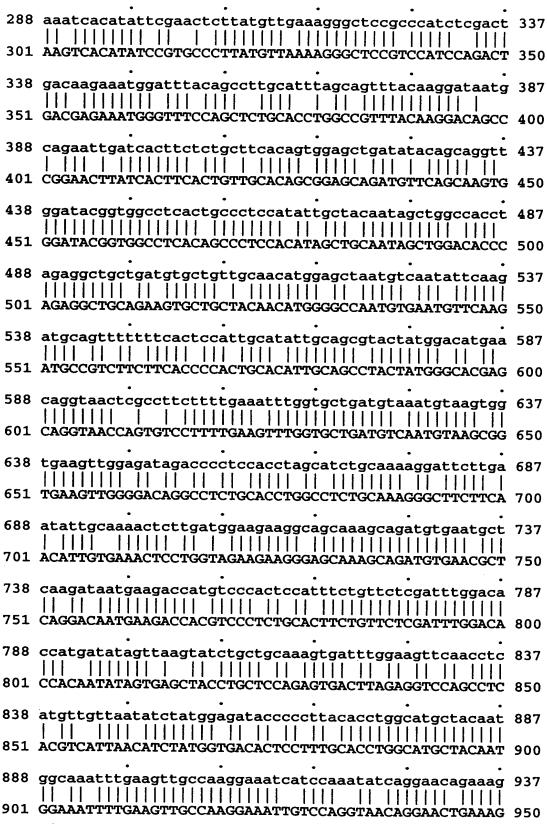
Fig. 7A

Serial No.: N/A Filed: Herewith
Title: Novel Cark Protein and Nucleic Acid Molecules and Uses

Therefor

Atty/Agent: Jean M. Silveri

Attorney Docket No.: MPI98-105P1RCP2DV1M

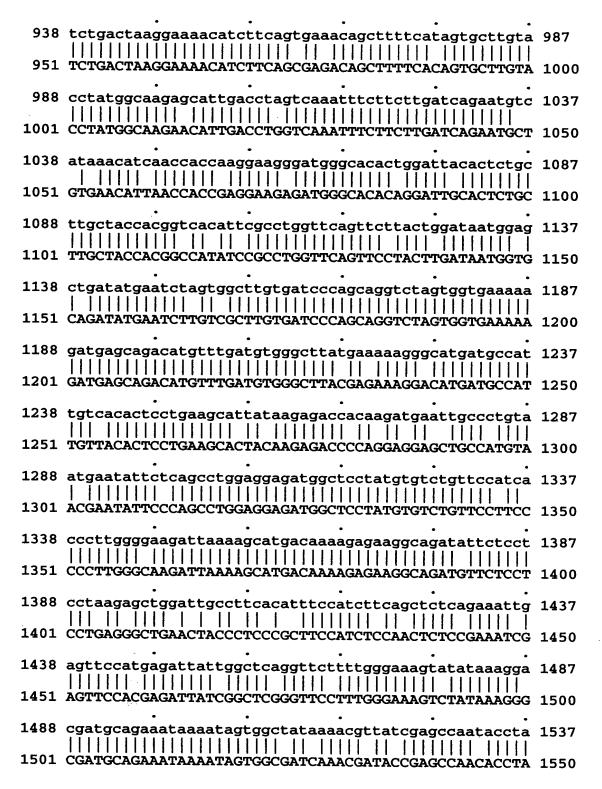


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Attorney Docket No.: MPI98-105P1RCP2DV1M



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Atty/Agent: Jean M. Silveri

Attorney Docket No.: MPI98-105P1RCP2DV1M

1538	ctgctccaagtcagatgtggatatgttttgccgagaggtgtccattctct	1587
1551	CTGCTCCAAGTCAGACGTGGATATGTTTTGCCGAGAGGTGTCCATTCTCT	1600
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1638	<pre>gatcccagccagtttgccattgtcactcaatacatatcagggggttctct </pre>	1687
1651	GÁCCCCÁGTCÁGTTTGCCÁTTGTCÁCTCÁGTÁCÁTTTCÁGGAGGCTCCCT	1700
	gttctccctccttcatgagcagaagaggattcttgatttgcagtctaaat	1737
	GTTCTCCCTGCTTCATGAACAGAAGAGAATTCTTGACTTGCAGTCTAAAT	1750
1738		1787
1788	TAATCATTGCGGTAGACGTTGCCAAGGGCATGGAGTACCTGCACAGCTTG	1800 1837
		1850
1838	tgaggatgggcatgctgtggtggcagattttggagaatcaagatttctac	1887
1851		1900
1888	agtctctggatgaagacaacatgacaaaacatgggaacctccgttgg	1937
1901		1950
1938	atggctcctgaggtgttcacgcagtgcactcggtacaccatcaaagcaga	1987
1951		2000
1988	tgtcttcagctatgctctgtgtctgtgggaaattctcactggcgaaattc	2037
	TGTCTTCAGTTACTCCCTGTGTCTGTGGGAGCTCCTCACTGGAGAAATTC	
	cattcgctcatctcaagccagcggctgcggcagcagacatggcttaccac	
	CATTCGCTCATCTCAAGCCAGCCAGCAGCAGATATGGCGTATCAC	
	cacatcagacctcccattggctattccattcccaagcccatatcatctct	
	CACATCAGACCGCCCATCGGCTATTCCATCCCCAAGCCCATCTCATCCCT	
	gctgatacgagggtggaacgcatgtcctgaaggaagacccgaattttctg	
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Serial No.: N/A Filed: Herewith
Title: Novel Cark Protein and Nucleic Acid Molecules and Uses

Therefor

Atty/Agent: Jean M. Silveri

Attorney Docket No.: MPI98-105P1RCP2DV1M

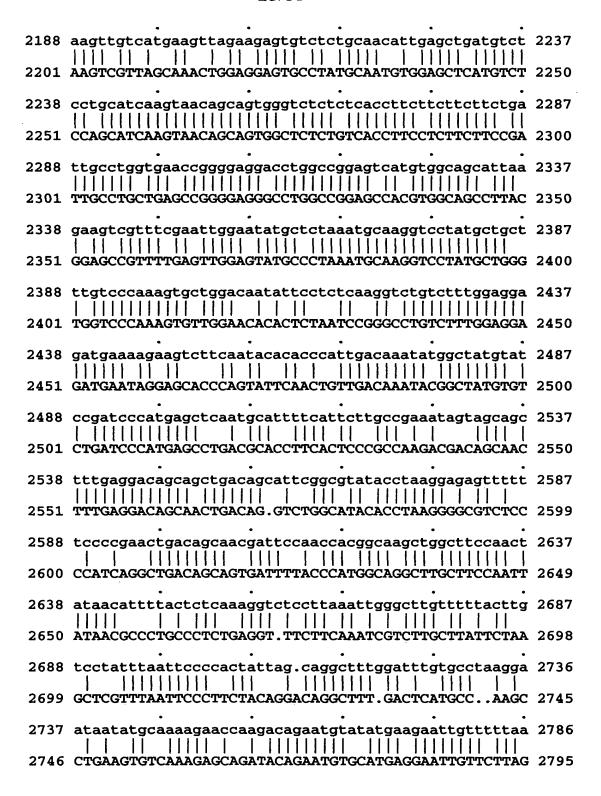


Fig. 7E

Serial No.: N/A Filed: Herewith
Title: Novel Cark Protein and Nucleic Acid Molecules and Uses

Therefor

Atty/Agent: Jean M. Silveri

Attorney Docket No.: MPI98-105P1RCP2DV1M

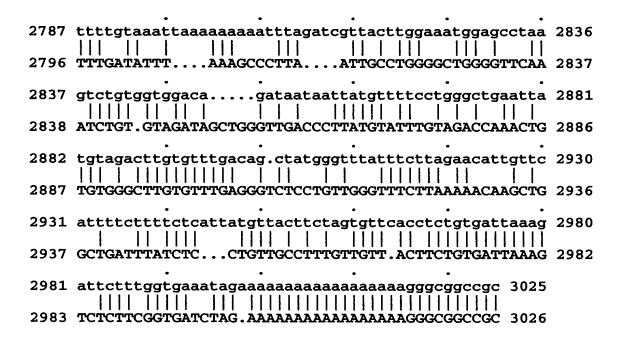


Fig. 7F

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Atty/Agent: Jean M. Silveri
        Attorney Docket No.: MPI98-105P1RCP2DV1M
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ratCARKpro (analysis only) - Import - complete
to: JAAa006_n check: 4709 from: 1 to: 835
carkprot (analysis only) - Import - complete
Symbol comparison table: /prod/ddm/seqanal/B:AST/matrix/aa/BLOSUM62
CompCheck: 1102
 Matrix made by matblas from blosum62.iij
      Gap Weight:
                    12
                          Average Match: 2.778
    Length Weight:
                        Average Mismatch: -2.248
                                         836
         Quality:
                  4079
                                Length:
           Ratio: 4.885
                                  Gaps:
Percent Similarity: 93.174
                        Percent Identity: 91.377
      Match display thresholds for the alignment(s):
                 = IDENTITY
                     2
IAAaoo6_n \times JAAa006_n
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                                                   rat
   1 MGNYKSRPTQTCTDEWKKKVSESYVITIERLEDDLQIKEKELTELRNIFG 50
                                                   human
51 SDEAFSEVSLNYRTERGLSLLHLCCVCGGNKSHIRALMLKGLRPSRLTRN 100
   51 SDEAFSKVNLNYRTENGLSLLHLCCICGGKKSHIRTLMLKGLRPSRLTRN 100
101 GFPALHLAVYKDSPELITSLLHSGADVQQVGYGGLTALHIAAIAGHPEAA 150
     101 GFTALHLAVYKDNAELITSLLHSGADIQQVGYGGLTALHIATIAGHLEAA 150
151 EVLLQHGANVNVQDAVFFTPLHIAAYYGHEQVTSVLLKFGADVNVSGEVG 200
   151 DVLLQHGANVNIQDAVFFTPLHIAAYYGHEQVTRLLLKFGADVNVSGEVG 200
201 DRPLHLASAKGFFNIVKLLVEEGSKADVNAQDNEDHVPLHFCSRFGHHNI 250
   201 DRPLHLASAKGFLNIAKLLMEEGSKADVNAQDNEDHVPLHFCSRFGHHDI 250
251 VSYLLQSDLEVQPHVINIYGDTPLHLACYNGNFEVAKEIVQVTGTESLTK 300
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Title: Novel Cark Protein and Nucleic Acid Molecules and Uses

Applicants: Jeyaseelan Raju

Serial No.: N/A

Therefor

Fig. 8A

251 VKYLLQSDLEVQPHVVNIYGDTPLHLACYNGKFEVAKEIIQISGTESLTK 300

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Title: Novel Cark Protein and Nucleic Acid Molecules and Uses

Therefor

Atty/Agent: Jean M. Silveri

Attorney Docket No.: MPI98-105P1RCP2DV1M

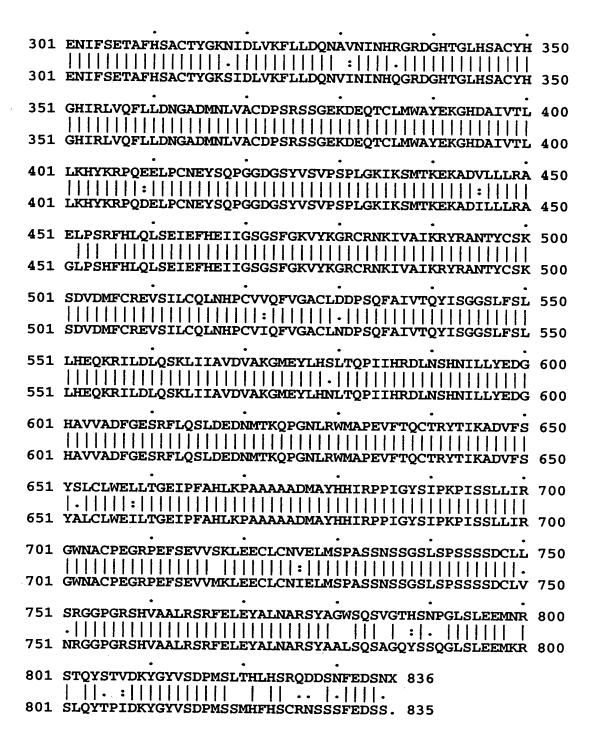


Fig. 8B

NIC **hCARK EXPRESSION MORMAL RA NORMAL LV** NORMAL LV **NORMAL LV** *HEART HEART* **HEART** ISCHEWIC IN **ISCHEWIC IN ISCHEWIC IV ISCHEWIC IN ISCHEWIC IN ISCHEWIC IN** ISCHEMIC HEART **HCM LA** FAMILIAL HCM RV 200.00 0.00

RELATIVE EXPRESSION (NTC AS REFERENCE)

Atty/Agent: Jean M. Silveri Attorney Docket No.: MPI98-105P1RCP2DV1M 32/35

Serial No.: N/A Title: Novel Cark Protein and Nucleic Acid Molecules and Uses

Therefor

Applicants: Jeyaseelan Raju

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Attorney Docket No.: MPI98-105P1RCP2DV1M

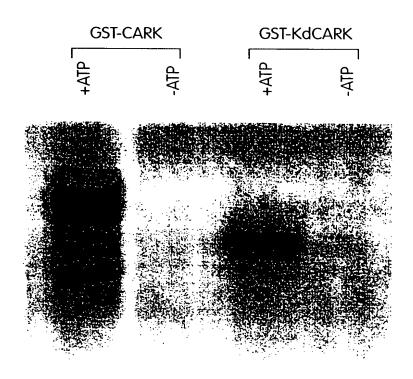


Fig. 10

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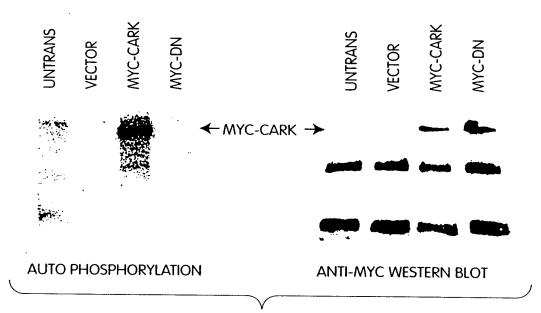


Fig. 11

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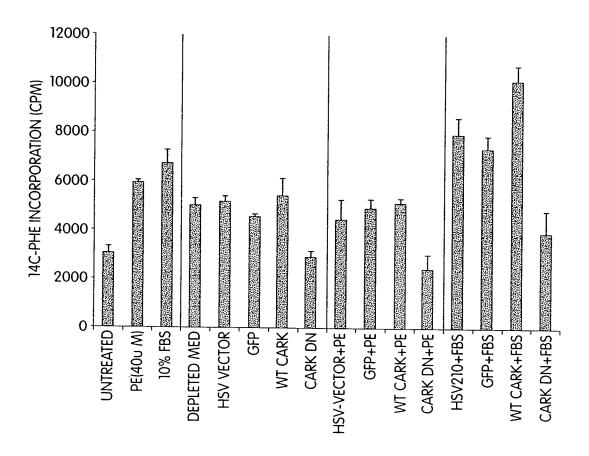


Fig. 12